

08/31/9, 831A

## Notice of Availability

### Applicant Aid for Biotechnology Computer Readable Form (CRF) Sequence Listings Submissions

The Patent and Trademark Office (PTO) has developed a computer program, called Checker, that will aid applicants in identifying and correcting errors prior to making submissions for compliance with the Requirements for Patent Applications Containing Nucleotide Sequence and/or Amino Acid Sequence Disclosures (sequence rules: 37 CFR 1.821 through 1.825). (Final rules were published in the *Federal Register* (55 FR 18230) on May 1, 1990, and in the *PTO Official Gazette* (1114 Off.Gaz.PatOffice 29) on May 15, 1990.)

Checker is a DOS-based software program that is intended to assist users in determining whether errors may be present in the sequence listings, and is not intended to guarantee that the submission is error-free.

The most current version of the software will be available via computer downloading (details below). Copies on diskette are also available. Updated software versions will not be automatically mailed out; any updates will be announced in the *PTO Official Gazette*.

The software can be accessed/requested in the following locations:

- 1) Dial-up access to the Patent and Trademark Office Bulletin Board System.  
Phone number: 703-305-8950  
Cost: Free-of-charge
- 2) Dial-up access through the Internet. FTP site: ftp.uspto.gov  
Login as "anonymous". Software is in directory /pub/checker  
Cost: Free-of-charge
- 3) For diskette copies, telephone requests to 703-306-2600.  
Cost: \$25.00

For Further Information Contact: Meredith Beckhardt at 703-308-4212.

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:36

INPUT SET: S9275.raw

This Raw Listing contains the General  
Information Section and those Sequences  
containing ERRORS.

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41

SEQUENCE LISTING

(1) General Information:

(i) APPLICANT: Hewick, Rodney M.  
Wang, Jack H.  
Wozney, John M.  
Celeste, Anthony J.

(ii) TITLE OF INVENTION: BONE AND CARTILAGE INDUCTIVE PROTEINS

(iii) NUMBER OF SEQUENCES: 44

(iv) CORRESPONDENCE ADDRESS:  
(A) ADDRESSEE: Legal Affairs - Genetics Institute, Inc.  
(B) STREET: 87 CambridgePark Drive  
(C) CITY: Cambridge  
(D) STATE: MA  
(E) COUNTRY: USA  
(F) ZIP: 02140

(v) COMPUTER READABLE FORM:  
(A) MEDIUM TYPE: Floppy disk  
(B) COMPUTER: IBM PC compatible  
(C) OPERATING SYSTEM: PC-DOS/MS-DOS  
(D) SOFTWARE: PatentIn Release #1.0, Version #1.30

(vi) CURRENT APPLICATION DATA:  
(A) APPLICATION NUMBER: US 08/319,831  
(B) FILING DATE: 06-OCT-1994  
(C) CLASSIFICATION:

(viii) ATTORNEY/AGENT INFORMATION:  
(A) NAME: Kapinos, Ellen J.  
(B) REGISTRATION NUMBER: 32,245  
(C) REFERENCE/DOCKET NUMBER: GI 5182A-DIV

(ix) TELECOMMUNICATION INFORMATION:  
(A) TELEPHONE: (617) 498-8622  
(B) TELEFAX: (617) 876-5851

Does Not Comply  
Corrected Diskette Needed

ERRORED SEQUENCES FOLLOW:

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:38

INPUT SET: S9275.raw

144 (2) INFORMATION FOR SEQ ID NO:5:  
145  
146 (i) SEQUENCE CHARACTERISTICS:  
--> 147 (A) LENGTH: 80 base pairs  
148 (B) TYPE: nucleic acid  
149 (C) STRANDEDNESS: double  
150 (D) TOPOLOGY: linear  
151  
152 (ii) MOLECULE TYPE: DNA (genomic)  
153  
154 (iii) HYPOTHETICAL: NO  
155  
156 (iv) ANTI-SENSE: NO  
157  
158 (vi) ORIGINAL SOURCE:  
159 (A) ORGANISM: Bos taurus  
160  
161 (vii) IMMEDIATE SOURCE:  
162 (B) CLONE: acc30  
163  
164 (viii) POSITION IN GENOME:  
165 (C) UNITS: bp  
166  
167 (ix) FEATURE:  
168 (A) NAME/KEY: CDS  
169 (B) LOCATION: 25..57  
170  
171 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:  
172  
173 GGATCCGCGT GCTGTGCTCC GACC AAG CTG AGC GCC ACC TCC GTG CTC TAC  
174 51  
175 Lys Leu Ser Ala Thr Ser Val Leu Tyr  
176 1 5  
177  
--> 178 TAC GAC AGCAGCAACA ATGTAATTCT AGA 80  
179 Tyr Asp  
180 10  
181

*move up to this line*

196 (2) INFORMATION FOR SEQ ID NO:7:  
197  
198 (i) SEQUENCE CHARACTERISTICS:  
--> 199 (A) LENGTH: 199 base pairs  
200 (B) TYPE: nucleic acid  
201 (C) STRANDEDNESS: double  
202 (D) TOPOLOGY: linear  
203  
204 (ii) MOLECULE TYPE: DNA (genomic)  
205  
206 (iii) HYPOTHETICAL: NO  
207  
208 (vi) ORIGINAL SOURCE:  
209 (A) ORGANISM: Bos taurus

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:40

INPUT SET: S9275.raw

210  
211 (vii) IMMEDIATE SOURCE:  
212 (A) LIBRARY: Bovine genomic  
213 (B) CLONE: Lambda 9800-10  
214  
215 (viii) POSITION IN GENOME:  
216 (C) UNITS: bp  
217  
218 (ix) FEATURE:  
219 (A) NAME/KEY: exon  
220 (B) LOCATION: 30..199  
221  
222 (ix) FEATURE:  
223 (A) NAME/KEY: intron  
224 (B) LOCATION: 1..29  
225  
226 (ix) FEATURE:  
227 (A) NAME/KEY: CDS  
228 (B) LOCATION: 30..179  
229  
230 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

--> 231  
232 TGCCCCGCTGC CCCCTCCCGC CCCCGCCAG GTG CAC CTG CTG AAG CCG CAC  
233 GCG 53  
234  
235  
236  
237 GTC CCC AAG GCG TGC TGC GCG CCC ACC AAG CTG AGC GCC ACT TCC GTG  
238 101  
239 Val Pro Lys Ala Cys Cys Ala Pro Thr Lys Leu Ser Ala Thr Ser Val  
240 10 15 20 25  
241  
242 CTC TAC TAC GAC AGC AGC AAC AAC GTC ATC CTG CGC AAG CAC CGC AAC  
243 149  
244 Leu Tyr Tyr Asp Ser Ser Asn Asn Val Ile Leu Arg Lys His Arg Asn  
245 30 35 40  
246  
247 ATG GTG GTC CGC GCC TGC GGC TGC CAC TGA GGCCCCAACT CCACCGGCAG  
248 199  
249 Met Val Val Arg Ala Cys Gly Cys His  
250 45 50  
251

once this total is  
moved up one line,  
amino acids should  
be aligned under codons  
Val His Leu Leu Lys  
1

---

252 (2) INFORMATION FOR SEQ ID NO:8:  
253

--> 254 (i) SEQUENCE CHARACTERISTICS:  
255 (A) LENGTH: 50 amino acids  
256 (B) TYPE: amino acid  
257 (D) TOPOLOGY: linear  
258

259 (ii) MOLECULE TYPE: protein

260  
261 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

49 are shown - is one  
missing?

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:43

INPUT SET: S9275.raw

262  
263 Val His Leu Leu Lys Pro His Ala Val Pro Lys Ala Cys Cys Ala Pro  
264 1 5 10 15  
265  
266 Thr Lys Leu Ser Ala Thr Ser Val Leu Tyr Tyr Asp Ser Ser Asn Asn  
267 20 25 30  
268  
269 Val Ile Leu Arg Lys His Arg Asn Met Val Val Arg Ala Cys Gly Cys  
270 35 40 45  
271  
272 His  
273  
274  
275  
276

---

277 (2) INFORMATION FOR SEQ ID NO:9:  
278  
279 (i) SEQUENCE CHARACTERISTICS:  
--> 280 (A) LENGTH: 172 base pairs  
281 (B) TYPE: nucleic acid  
282 (C) STRANDEDNESS: double  
283 (D) TOPOLOGY: linear  
284  
285 (ii) MOLECULE TYPE: DNA (genomic)  
286  
287 (iii) HYPOTHETICAL: NO  
288  
289 (vi) ORIGINAL SOURCE:  
290 (A) ORGANISM: Bos taurus  
291  
292 (vii) IMMEDIATE SOURCE:  
293 (A) LIBRARY: Bovine genomic  
294 (B) CLONE: Lambda 9800-10  
295  
296 (viii) POSITION IN GENOME:  
297 (C) UNITS: bp  
298  
299 (ix) FEATURE:  
300 (A) NAME/KEY: exon  
301 (B) LOCATION: 51..161  
302  
303 (ix) FEATURE:  
304 (A) NAME/KEY: intron  
305 (B) LOCATION: 1..50  
306  
307 (ix) FEATURE:  
308 (A) NAME/KEY: intron  
309 (B) LOCATION: 162..172  
310  
311 (ix) FEATURE:  
312 (A) NAME/KEY: CDS  
313 (B) LOCATION: 51..161

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:45

INPUT SET: S9275.raw

314  
315 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:9:  
316  
317 GGGGTGGGAG GGCACGTGGA TGGGACTCAC CTTCTCCCAC TACCCCCCAG GAC  
--> 318 TGG 56  
319 Asp Trp  
320 1  
321 GTC ATC GCC CCC CAA GGC TAC TCA GCC TAT TAC TGT GAA GGG GAG TGC  
322 104  
323 Val Ile Ala Pro Gln Gly Tyr Ser Ala Tyr Tyr Cys Glu Gly Glu Cys  
324 5 10 15  
325  
326 TCC TTC CCG CTG GAC TCC TGC ATG AAC GCC ACC AAC CAC GCC ATC CTG  
327 152  
328 Ser Phe Pro Leu Asp Ser Cys Met Asn Ala Thr Asn His Ala Ile Leu  
329 20 25 30  
330  
--> 331 CAG TCC CTG GTCAGTACCT C 172  
332 Gln Ser Leu  
333 5  
334

---

355 (2) INFORMATION FOR SEQ ID NO:11:  
356  
357 (i) SEQUENCE CHARACTERISTICS:  
--> 358 (A) LENGTH: 119 base pairs  
359 (B) TYPE: nucleic acid  
360 (C) STRANDEDNESS: double  
361 (D) TOPOLOGY: linear  
362  
363 (ii) MOLECULE TYPE: DNA (genomic)  
364  
365 (iii) HYPOTHETICAL: NO  
366  
367 (vi) ORIGINAL SOURCE:  
368 (A) ORGANISM: Bos taurus  
369  
370 (vii) IMMEDIATE SOURCE:  
371 (A) LIBRARY: Bovine genous  
372 (B) CLONE: Lambda 9800-10  
373  
374 (viii) POSITION IN GENOME:  
375 (C) UNITS: bp  
376  
377 (ix) FEATURE:  
378 (A) NAME/KEY: exon  
379 (B) LOCATION: 20..99  
380  
381 (ix) FEATURE:  
382 (A) NAME/KEY: intron  
383 (B) LOCATION: 1..19  
384  
385 (ix) FEATURE:

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:47

INPUT SET: S9275.raw

386 (A) NAME/KEY: intron  
387 (B) LOCATION: 100..119  
388  
389 (ix) FEATURE:  
390 (A) NAME/KEY: CDS  
391 (B) LOCATION: 22..99  
392  
393  
394 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:  
395  
396 CCCTTGCGTG TCCCCGCAGA C GAC GTC CAC GGC TCC CAC GGC CGG CAG  
--> 397 GTG 51  
398 Asp Val His Gly Ser His Gly Arg Gln Val  
399 1 5 10  
400  
401 TGC CGT CGG CAC GAG CTG TAC GTG AGC TTC CAG GAC CTG GGC TGG CTG  
402 99  
403 Cys Arg Arg His Glu Leu Tyr Val Ser Phe Gln Asp Leu Gly Trp Leu  
404 15 20 25  
405  
--> 406 GTGAGTTCCG ACTCTCCTTT  
407

---

425 (2) INFORMATION FOR SEQ ID NO:13:  
426  
427 (i) SEQUENCE CHARACTERISTICS:  
--> 428 (A) LENGTH: 1003 base pairs  
429 (B) TYPE: nucleic acid  
430 (C) STRANDEDNESS: double  
431 (D) TOPOLOGY: circular  
432  
433 (ii) MOLECULE TYPE: cDNA to mRNA  
434  
435 (iii) HYPOTHETICAL: NO  
436  
437 (vi) ORIGINAL SOURCE:  
438 (A) ORGANISM: Homo sapiens  
439 (F) TISSUE TYPE: Human Heart  
440  
441 (vii) IMMEDIATE SOURCE:  
442 (A) LIBRARY: Human heart cDNA library stratagene catalog  
443 (B) CLONE: hh38  
444  
445 (viii) POSITION IN GENOME:  
446 (C) UNITS: bp  
447  
448 (ix) FEATURE:  
449 (A) NAME/KEY: CDS  
450 (B) LOCATION: 8..850  
451  
452 (ix) FEATURE:  
453 (A) NAME/KEY: mat\_peptide  
454 (B) LOCATION: 427..843

RAW SEQUENCE LISTING  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:50

INPUT SET: S9275.raw

455

456

(ix) FEATURE:

457

(A) NAME/KEY: mRNA

458

(B) LOCATION: 1..997

459

460

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:

461

462

GAATTCC GAG CCC CAT TGG AAG GAG TTC CGC TTT GAC CTG ACC CAG ATC

463

49

464

Glu Pro His Trp Lys Glu Phe Arg Phe Asp Leu Thr Gln Ile

465

-139

-135

-130

466

467

CCG GCT GGG GAG GCG GTC ACA GCT GCG GAG TTC CGG ATT TAC AAG GTG

468

97

469

Pro Ala Gly Glu Ala Val Thr Ala Ala Glu Phe Arg Ile Tyr Lys Val

470

-125

-120

-115

-110

471

472

CCC AGC ATC CAC CTG CTC AAC AGG ACC CTC CAC GTC AGC ATG TTC CAG

473

145

474

Pro Ser Ile His Leu Leu Asn Arg Thr Leu His Val Ser Met Phe Gln

475

-105

-100

-95

476

477

GTG GTC CAG GAG CAG TCC AAC AGG GAG TCT GAC TTG TTC TTT TTG GAT

478

193

479

Val Val Gln Glu Gln Ser Asn Arg Glu Ser Asp Leu Phe Phe Leu Asp

480

-90

-85

-80

481

482

CTT CAG ACG CTC CGA GCT GGA GAC GAG GGC TGG CTG GTG CTG GAT GTC

483

241

484

Leu Gln Thr Leu Arg Ala Gly Asp Glu Gly Trp Leu Val Leu Asp Val

485

-75

-70

-65

486

487

ACA GCA GCC AGT GAC TGC TGG TTG CTG AAG CGT CAC AAG GAC CTG GGA

488

289

489

Thr Ala Ala Ser Asp Cys Trp Leu Leu Lys Arg His Lys Asp Leu Gly

490

-60

-55

-50

491

492

CTC CGC CTC TAT GTG GAG ACT GAG GAT GGG CAC AGC GTG GAT CCT GGC

493

337

494

Leu Arg Leu Tyr Val Glu Thr Glu Asp Gly His Ser Val Asp Pro Gly

495

-45

-40

-35

-30

496

497

CTG GCC GGC CTG CTG GGT CAA CGG GCC CCA CGC TCC CAA CAG CCT TTC

498

385

499

Leu Ala Gly Leu Leu Gly Gln Arg Ala Pro Arg Ser Gln Gln Pro Phe

500

-25

-20

-15

501

502

GTG GTC ACT TTC TTC AGG GCC AGT CCG AGT CCC ATC CGC ACC CCT CGG

503

433

504

Val Val Thr Phe Phe Arg Ala Ser Pro Ser Pro Ile Arg Thr Pro Arg

505

-10

-5

1

506

507

*edit  
throughout*



# RAW SEQUENCE LISTING PATENT APPLICATION US/08/319,831A

DATE: 05/09/96  
TIME: 16:30:52

INPUT SET: S9275.raw

```

508 GCA GTG AGG CCA CTG AGG AGG AGG CAG CCG AAG AAA AGC AAC GAG CTG
509 481
510 Ala Val Arg Pro Leu Arg Arg Arg Gln Pro Lys Lys Ser Asn Glu Leu
511      5              10              15
512
513 CCG CAG GCC AAC CGA CTC CCA GGG ATC TTT GAT GAC GTC CAC GGC TCC
514 529
515 Pro Gln Ala Asn Arg Leu Pro Gly Ile Phe Asp Asp Val His Gly Ser
516 20              25              30              35
517
518 CAC GGC CGG CAG GTC TGC CGT CGG CAC GAG CTC TAC GTC AGC TTC CAG
519 577
520 His Gly Arg Gln Val Cys Arg Arg His Glu Leu Tyr Val Ser Phe Gln
521      40              45              50
522
523 GAC CTT GGC TGG CTG GAC TGG GTC ATC GCC CCC CAA GGC TAC TCA GCC
524 625
525 Asp Leu Gly Trp Leu Asp Trp Val Ile Ala Pro Gln Gly Tyr Ser Ala
526      55              60              65
527
528 TAT TAC TGT GAG GGG GAG TGC TCC TTC CCG CTG GAC TCC TGC ATG AAC
529 673
530 Tyr Tyr Cys Glu Gly Glu Cys Ser Phe Pro Leu Asp Ser Cys Met Asn
531      70              75              80
532
533 GCC ACC AAC CAC GCC ATC CTG CAG TCC CTG GTG CAC CTG ATG AAG CCA
534 721
535 Ala Thr Asn His Ala Ile Leu Gln Ser Leu Val His Leu Met Lys Pro
536      85              90              95
537
538 AAC GCA GTC CCC AAG GCG TGC TGT GCA CCC ACC AAG CTG AGC GCC ACC
539 769
540 Asn Ala Val Pro Lys Ala Cys Cys Ala Pro Thr Lys Leu Ser Ala Thr
541 100              105              110              115
542
543 TCT GTG CTC TAC TAT GAC AGC AGC AAC AAC GTC ATC CTG CGC AAG CAC
544 817
545 Ser Val Leu Tyr Tyr Asp Ser Ser Asn Asn Val Ile Leu Arg Lys His
546      120              125              130
547
548 CGC AAC ATG GTG GTC AAG GCC TGC GGC TGC CAC
--> 549 TGAGTCAGCCGCCCCAGCCC 870
550 Arg Asn Met Val Val Lys Ala Cys Gly Cys His
551      135              140
552
553 TACTGCAGCC ACCCTTCTCA TCTGGATCGG GCCCTGCAGA
--> 554 GGCAGAAAACCTTAAATGC 930
555
556 TGTCACAGCT CAAGCAGGAG TGTCAGGGGC CCTCACTCTC
--> 557 GGTGCCTACTTCCTGTCAGG 990
558
559 CTTCTGGGAA TTC
560

```

base must be  
in group  
of 10  
per 1.822 (f)  
of sequence  
Rules

SEQUENCE VERIFICATION REPORT  
PATENT APPLICATION US/08/319,831ADATE: 05/09/96  
TIME: 16:30:54

INPUT SET: S9275.raw

Line	Error	Original Text
147	Entered (80) and Calc. Seq. Length (29) differ	(A) LENGTH: 80 base pairs
178	# of Sequences for line conflicts w/ running total	TAC GAC AGCAGCAACA ATGTAATTCT AGA
199	Entered (199) and Calc. Seq. Length (3) differ	(A) LENGTH: 199 base pairs
233	# of Sequences for line conflicts w/ running total	GCG 53
255	Entered (50) and Calc. Seq. Length (49) differ	(A) LENGTH: 50 amino acids
280	Entered (172) and Calc. Seq. Length (23) differ	(A) LENGTH: 172 base pairs
318	# of Sequences for line conflicts w/ running total	TGG 56
331	# of Sequences for line conflicts w/ running total	CAG TCC CTG GTCAGTACCT C
358	Entered (119) and Calc. Seq. Length (23) differ	(A) LENGTH: 119 base pairs
397	# of Sequences for line conflicts w/ running total	GTG 51
406	# of Sequences for line conflicts w/ running total	GTGAGTTCCG ACTCTCCTTT
428	Entered (1003) and Calc. Seq. Length (60) differ	(A) LENGTH: 1003 base pairs
549	# of Sequences for line conflicts w/ running total	TGAGTCAGCCCGCCCAGCCC 870
554	# of Sequences for line conflicts w/ running total	GGCAGAAAACCCTTAAATGC 930
557	# of Sequences for line conflicts w/ running total	GGTGCCTACTTCCTGTCAGG 990